CASWELL FILE 334B

UNITED STATES ENVIRONMENTAL PROTECTION AGENCY
WASHINGTON, D.C. 20460

JAN 11 1994

OPP OFFICIAL RECORD
HEALTH EFFECTS DIVISION
SCIENTIFIC DATA REVIEWS
EPA SERIES 361

OFFICE OF
PREVENTION, PESTICIDES AND
TOXIC SUBSTANCES

**MEMORANDUM**

pc code 108102

SUBJECT: I.D. 108102-010182: Pirimiphos Methyl, Data for 81-8ss and 82-7ss neurotoxicity studies.

FROM: Marion Copley, DVM, Section Head
Section 4, Tox. Br. 1
Health Effects Division (7509C)

Marion Copley 1/11/94

TO: Myers/Kendall (PM # 51)
Special Review and Reregistration Division (7508W)

AND: Edwards (PM # 12)
Registration Division (7505C)

THRU: John Doherty, PhD 1/11/94
Section 4, Tox. Br. 1
Health Effects Division (7509C)

DP Barcode #: D196638
Submission #: S453100
Tox. Chem. #: 334B
P.C.Code: 108102

**CONCLUSIONS:**

The submitted hen delayed type neurotoxicity studies and the rat subchronic toxicity study do not satisfy the guideline requirements for mammalian acute (81-8ss) and subchronic (82-7ss) neurotoxicity screening studies. Rats must be used for the neurotoxicity screen studies and the existing rat study did not include the special assessments for FOB or motor activity. These study types remain data gaps. It is recommended that the protocol for the series 82-7ss study be submitted to the Agency for review.

**ACTION REQUESTED:**

Zeneca, Inc. (see letter from Compliance Services International to Ron Kendall, RD, dated 10/27/93) has requested that the Agency use the studies listed below to satisfy the requirements for the Acute Neurotoxicity Screen and Rat (81-8ss), 90-day Neurotoxicity Screen, Rat (82-7ss). These studies were submitted previously by ICI Americas, Inc. to satisfy the above requirements.

cc: Chow, CCB

**DISCUSSION:**

**90-day Oral Toxicity in Rats: Report No. HO/IH/R/284 (1970) (MRID # 80745)**

**Determination of No Effect Level During a 28 Day Rat Feeding Study: Report No. CTL/P/199 (1975) (MRID #1259343)**

The above two studies with rats cannot be used to satisfy the series 81-8ss and 82-7ss special neurotoxicity screens because they did not include the specific FOB, motor activity or pathological assessments now required. Their protocols and submission formats are also otherwise not consistent with the current requirements. The registrant is advised to consult the Federal Register for the specific requirements for series 81-8ss and 82-7ss neurotoxicity screen studies.

**Examination of Pirimiphos Methyl for Neurotoxicity in the Domestic Hen: ICI/49/75220 (1978) (MRID #80721)**

**Acute Delayed Neurotoxicity Study in the Domestic Hen:Lab Project Number: CTL/C/2167 (1989) (MRID # 41599503)**

**The Acute Oral Toxicity (LD50) and Neurotoxic Effects of Pirimiphos-Methyl on the Domestic Hen: Report No. not given (1983) (MRID #42986101)**

These studies with <u>hens</u> cannot be used to satisfy the series 81-8ss and 82-7ss neurotoxicity screen studies which must be conducted with <u>rats.</u>

Note: The data requirement for the series 81-7 acute delayed neurotoxicity study with hens is not addressed in the above. The phase IV review of data submitted by the ICI company (former name of Zeneca) indicated that there are data for this series that are adequate for further review. The final determination of the acceptability of these data to satisfy the series 81-7 requirement will be made in Phase V review.

\NDOHERTY\D196638

13544

# 009782

| | |
|---|---|
| **Chemical:** | **Pirimiphos-methyl (ANSI)** |
| **PC Code:** | **108102** |
| **HED File Code** | ***13000 Tox Reviews*** |
| **Memo Date:** | **01/11/94** |
| **File ID:** | **DPD196638** |
| **Accession Number:** | **412-02-0005** |

**HED Records Reference Center**
**10/25/2001**